

```
ring nodes :
    1 2 3 4 5 6 10

chain bonds :
    2-7 5-15 7-8 7-9 7-10 10-11 11-12 11-13 13-14

ring bonds :
    1-2 1-6 2-3 3-4 4-5 5-6

exact/norm bonds :
    1-2 1-6 2-3 2-7 3-4 4-5 5-6 7-8 7-9 7-10 11-12 11-13 13-14

exact bonds :
    5-15 10-11

isolated ring systems :
    containing 1 :
```

7 8 9 11 12 13 14 15

```
Match level:
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 9:CLASS 10:Atom 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS
```

=> s l1

SAMPLE SEARCH INITIATED 11:57:57 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 68 TO ITERATE

100.0% PROCESSED 68 ITERATIONS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

PROJECTED ITERATIONS: 866 TO 1854
PROJECTED ANSWERS: 2 TO 124

L2 2 SEA SSS SAM L1

=> s l1 sss full

FULL SEARCH INITIATED 11:58:21 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 1413 TO ITERATE

100.0% PROCESSED 1413 ITERATIONS 103 ANSWERS

2 ANSWERS

SEARCH TIME: 00.00.01

L3 103 SEA SSS FUL L1

=> file caplus

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION

FULL ESTIMATED COST 161.76 161.97

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=> s 13

L4 4 L3

=> d l4 1-4 bib abs hitstr

L4 ANSWER 1 OF 4 CAPLUS COPYRIGHT 2005 ACS on STN

AN 2005:1028086 CAPLUS

DN 143:326396

- Preparation of piperidinyl- and piperazinyl-sulfonylmethyl hydroxamic ΤI acids and their use as protease inhibitors
- Mcdonald, Joseph J.; Kassab, Darren J.; Massa, Mark A.; Grapperhaus, IN Margaret L.; Schmidt, Michelle A.; Rico, Joseph G.; Mullins, Patrick B.; Brown, David L.
- PA USA
- U.S. Pat. Appl. Publ., 417 pp., Cont.-in-part of U.S. Ser. No. 618,288. so CODEN: USXXCO
- DTPatent
- English LA

| NT 3 |
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| FAN.CNT 3 | | | | | | | | | | | | | | • | | | | | |
|-----------|-----|--------------|------|-----|-----------------------------|------|----------------|------|----------------|-----|-----------------|-----|----------|------|----------|-----|-------|-----|--|
| | PA' | CENT | NO. | | | KIN | | DATE | | | APPLICATION NO. | | | | | | DATE | | |
| PI | US | 2005 | | | | | US 2003-700202 | | | | | | 20031103 | | | | | | |
| | US | 2005 | 38 | | | | | | US 2003-618288 | | | | | | 20030425 | | | | |
| | WO | 2005 | 21 | A2 | A2 20050512 WO 2004-US36666 | | | | | | | | | | | | | | |
| | WO | 2005 | A3 | | 20050707 | | | | | | | | | 0041 | 103 | | | | |
| | | W: | | | | | | AU, | AZ, | BA, | BB, | BG, | BR. | BW. | BY. | BZ. | CA. | CH. | |
| | | | CN, | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DZ, | EC. | EE. | EG. | ES. | FI. | GB. | GD, | |
| | | | GE, | GH, | GM, | HR, | HU, | ID, | IL, | IN, | ıs, | JP. | KE. | KG. | KP. | KR. | KZ. | T.C | |
| | | | LK, | LR, | LS, | LT, | LU, | LV, | MA, | MD, | MG. | MK. | MN. | MW. | MX. | M2. | NA AN | NT | |
| | | | NO, | NZ, | OM, | PG, | PH, | PL, | PT, | RO, | RU, | sc. | SD. | SE. | SG. | SK. | SL. | SY | |
| | | | ТJ, | TM, | TN, | TR, | TT, | TZ, | UA, | UG, | US, | UZ. | VC, | VN. | YU. | ZA. | ZM. | 2.W | |
| | | RW: | BW, | GH, | GM, | KE, | LS, | MW, | MZ, | NA, | SD, | SL, | sz. | TZ. | UG. | ZM. | ZW. | AM. | |
| | | | ΑZ, | BY, | KG, | ΚZ, | MD, | RU, | ТJ, | TM, | AT, | BE, | BG, | CH, | CY. | CZ. | DE. | DK. | |
| | | | EE, | ES, | FI, | FR, | GB, | GR, | HU, | IE, | IS, | IT, | LU, | MC. | NL. | PL. | PT. | RO. | |
| | | | SE, | SI, | SK, | TR, | BF, | ВJ, | CF, | CG, | CI, | CM, | GA, | GN, | GO, | GW, | ML. | MR. | |
| | | | ΝE, | SN, | TD, | TG | | | | | - | - | • | • | ~, | - , | , | | |
| PRAI | US | 2002-375598P | | | | P | | 2002 | 0425 | | | | | | | | | | |
| | US | 2002-380713P | | | | P | | 2002 | 0515 | | | | | | | | | | |
| | US | 2002-392021P | | | | P | | 2002 | 0627 | | | | | | | | | | |
| | US | 2003 | -618 | 288 | | A2 | | 2003 | 0425 | | | | | | | | | | |
| | US | 2003 | Α | | 2003 | 1103 | | | | | | | | | | | | | |

Title compds. I [A1 and A2 together with the C to which they are bonded AB join to form (un) substituted heterocyclyl or carbocyclyl, or A1 and A2 are independently selected from H, alkyl, alkoxyalkyl, alkenyl, alkynyl, etc.; Rx = H, halo, CN, OH, NO2, alkyl, alkenyl, alkoxy, alkoxyalkyl,

heterocyclyl, etc.; Y = N, CH, or CRx; E1 = (un)substituted carbocyclyl, heterocyclyl, etc.; E2 = 0, C0, C(0)0, OC(0), bond, S, etc.; E3 = halo, CN, (un) substituted alkyl, alkenyl, alkynyl, heterocyclyl, heterocyclylalkyl, etc.] and their pharmaceutically acceptable salts are prepd. and disclosed as protease inhibitors. E.g., a multi-step synthesis of II, starting from Et crotyl phosphonate and tert-Bu 4-[(4-formylpiperidin-1-yl)sulfonyl]tetrahydro-2H-pyran-2H-pyran-4carboxylate, was given. This invention is directed generally to proteinase (also known as 'protease') inhibitors, and more particularly, inhibitors of matrix metalloproteinase (also known as 'matrix metalloprotease' or 'MMP'), aggrecanase, or TNF-.alpha. convertase activity. In assays to det. inhibition consts. (Ki) against MMP-1, MMP-2, MMP-9, MMP-13 and MMP-14, I possessed values ranging from 0.13->10,000. This invention also is directed to compns. of such hydroxamic acids, intermediates for the syntheses of such hydroxamic acids, methods for making such hydroxamic acids, and methods for treating conditions (particularly pathol. conditions) assocd. with MMP, aggrecanase, or TNF-.alpha. convertase activity. 622387-48-8P 622389-78-0P 622390-02-7P

1T 622387-48-8P 622389-78-0P 622390-02-7P 622390-05-0P 622390-07-2P 622390-55-0P 622390-62-9P 622390-68-5P 622390-72-1P 622390-78-7P 622390-81-2P 622390-88-9P 622391-15-5P 622391-31-5P 622391-35-9P 622391-39-3P 622391-39-3P 622391-39-7P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(intermediate; prepn. of piperidinyl- and piperazinyl-sulfonylmethyl hydroxamic acids and their use as matrix metalloproteinase inhibitors) 622387-48-8 CAPLUS

2H-Pyran-4-carboxamide, tetrahydro-4-[(4-octyl-1-piperidinyl)sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622389-78-0 CAPLUS
CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-[(phenylmethoxy)methyl]-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN

CN

RN 622390-02-7 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-[(2,2,3,3-tetrafluoropropoxy)methyl]-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622390-05-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[[[4-(trifluoromethyl)phenyl]methoxy]methyl]-1-piperidinyl]sulfonyl]-(9CI) (CA INDEX NAME)

RN 622390-07-2 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-(1Z)-1-octenyl-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622390-55-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[5-(2,2,2-trifluoroethoxy)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622390-62-9 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-(5-ethoxypentyl)-1-piperidinyl]sulfonyl]tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622390-68-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-(4-propoxybutyl)-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622390-72-1 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[4-(4,4,4-trifluorobutoxy)butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622390-78-7 CAPLUS
CN 2H-Pyran-4-carboxamide, 4-[[4-[(hexyloxy)methyl]-1piperidinyl]sulfonyl]tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI)
(CA INDEX NAME)

RN 622390-81-2 CAPLUS
CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[(4,4,4-trifluorobutoxy)methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622390-88-9 CAPLUS
CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-(6,6,6-trifluorohexyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622390-93-6 CAPLUS
CN 2H-Pyran-4-carboxamide, 4-[[4-(2-butoxyethyl)-1piperidinyl]sulfonyl]tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI)
(CA INDEX NAME)

RN 622390-99-2 CAPLUS
CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[2-(4,4,4-trifluorobutoxy)ethyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622391-09-7 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[4-(3,3,3-trifluoropropoxy)butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622391-15-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-[2-(pentyloxy)ethyl]-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622391-31-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-(7,7,8,8,8-pentafluorooctyl)-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622391-35-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-(8,8,8-trifluorooctyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622391-39-3 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-(5,5,6,6,6-pentafluorohexyl)-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622391-43-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-(6,6,7,7,7-pentafluoroheptyl)-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622391-83-7 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[3-[4-(trifluoromethoxy)phenyl]-1,2,4-oxadiazol-5-yl]butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

IT 622382-02-9P 622385-57-3P 622385-59-5P 622385-60-8P 622385-61-9P 622385-62-0P 622385-88-0P 622385-89-1P 622385-90-4P 622385-91-5P 622385-92-6P 622387-62-6P 622387-72-8P 622387-73-9P 622387-74-0P 622387-75-1P 622387-76-2P 622387-77-3P 622387-78-4P 622387-79-5P 622387-80-8P 622387-81-9P 622387-82-0P 622387-83-1P 622387-84-2P 622387-92-2P 622387-94-4P 622387-96-6P 622387-98-8P 622388-18-5P 622388-35-6P 622388-36-7P 622388-37-8P 622388-38-9P 622388-40-3P 622388-42-5P 622388-44-7P 622388-46-9P 622388-47-0P 622388-48-1P 622388-50-5P 622388-51-6P 622388-52-7P 622388-53-8P 622388-54-9P 622388-55-0P 622388-56-1P 622388-57-2P 622388-58-3P 622388-60-7P 622388-61-8P

CN

622388-62-9P 622388-63-0P 622388-67-4P 622388-72-1P 622388-75-4P 622388-76-5P 622388-79-8P 622389-05-3P 622389-06-4P 622389-10-0P 622389-11-1P 622389-12-2P 622389-24-6P 622389-27-9P 622389-73-5P 851771-04-5P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of piperidinyl- and piperazinyl-sulfonylmethyl hydroxamic acids and their use as matrix metalloproteinase inhibitors)

RN 622382-02-9 CAPLUS

2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[(4-octyl-1-piperidinyl)sulfonyl]- (9CI) (CA INDEX NAME)

RN 622385-57-3 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(phenylmethoxy)methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622385-59-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(2,2,3,3-tetrafluoropropoxy)methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_2CH-CF_2-CH_2-O-CH_2$$

$$0$$

$$0$$

$$C-NH-OH$$

$$0$$

RN 622385-60-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(2,2,3,3,3-pentafluoropropoxy)methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3C-CF_2-CH_2-O-CH_2$$
 $N-S$
 $C-NH-OH$
 O
 C

RN 622385-61-9 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(2,2,3,3,4,4,4-heptafluorobutoxy)methyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

$$F_3C-CF_2-CF_2-CH_2-O-CH_2$$
 $N-S$
 $C-NH-OH$
 O
 C

RN 622385-62-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[[(2,2,3,3,4,4,5,5-octafluoropentyl)oxy]methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_2CH-(CF_2)_3-CH_2-O-CH_2$$
 $N-S$
 0
 $C-NH-OH$
 0

RN 622385-88-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(1Z)-1-octenyl-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

Me
$$(CH_2)_5$$
 Z

RN 622385-89-1 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(1E)-1-octenyl-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622385-90-4 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[[[4-(trifluoromethyl)phenyl]methoxy]methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$CH_2-O-CH_2$$
 N
 CH_2-O-CH_2
 CH_2-O-CH_2
 CH_2-O-CH_2
 CH_2-O-CH_2

RN 622385-91-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[[[4-(trifluoromethoxy)phenyl]methoxy]methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3C-O$$
 CH_2-O-CH_2
 $N-S$
 $C-NH-OH$

RN 622385-92-6 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[[4-[[4-[(trifluoromethyl)thio]phenyl]methoxy]methyl]-1-piperidinyl]sulfonyl]-(9CI) (CA INDEX NAME)

RN 622387-62-6 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-(2,2,2-trifluoroethoxy)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-72-8 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-(3-butoxypropyl)-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

RN 622387-73-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[3-(4,4,4-trifluorobutoxy)propyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-74-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-(3,3,3-trifluoropropoxy)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-75-1 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-(5-ethoxypentyl)-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

RN 622387-76-2 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(4-propoxybutyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-77-3 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[4-(4,4,4-trifluorobutoxy)butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3C-(CH_2)_3-O-(CH_2)_4$$
 $N-S$
 $C-NH-OH$

RN 622387-78-4 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(hexyloxy)methyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

RN 622387-79-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(4,4,4-trifluorobutoxy)methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3C-(CH_2)_3-O-CH_2$$
 $N-S$
 $C-NH-OH$

RN 622387-80-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(6,6,6-trifluorohexyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-81-9 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-(2-butoxyethyl)-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

RN 622387-82-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[2-(4,4,4-trifluorobutoxy)ethyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-83-1 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[4-(3,3,3-trifluoropropoxy)butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-84-2 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[2-(pentyloxy)ethyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-92-2 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(7,7,8,8,8-pentafluorooctyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-94-4 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(8,8,8-trifluorooctyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-96-6 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(5,5,6,6,6-pentafluorohexyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-98-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(6,6,7,7,7-pentafluoroheptyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-18-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[4-[3-[4-(trifluoromethoxy)phenyl]-1,2,4-oxadiazol-5-yl]butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-35-6 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[5-(3,4-dihydro-2(1H)-isoquinolinyl)-5-oxopentyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy-(9CI) (CA INDEX NAME)

RN 622388-36-7 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(1E,3E)-5-(3,4-dihydro-2(1H)-isoquinolinyl)-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-37-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(1E,3E)-5-oxo-5-(4-phenyl-1-piperazinyl)-1,3-pentadienyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-38-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(1E,3E)-5-[methyl(2-methylpropyl)amino]-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-40-3 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(1E,3E)-5-oxo-5-(1-piperidinyl)-1,3-pentadienyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-42-5 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(1E,3E)-5-(cyclohexylmethylamino)-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-44-7 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(1E,3E)-5-(2,3-dihydro-1H-indol-1-yl)-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-46-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(1E,3E)-5-[(3aR,7aS)-octahydro-2H-isoindol-2-yl]-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]-, rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

Double bond geometry as shown.

RN 622388-47-0 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(1E,3E)-5-(1,3-dihydro-2H-isoindol-2-yl)-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-48-1 CAPLUS

CN 4-Piperidinepentanamide, N-methyl-N-2-propynyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} \text{Me O} & & & \\ & | & | \\ & | & | \\ \text{HC} = \text{C-CH}_2 - \text{N-C-} (\text{CH}_2)_4 & & & \\ & 0 & & | \\ & 0 & & | \\ & 0 & & | \\ \end{array}$$

RN -622388-50-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-(4-morpholinyl)-5-oxopentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-51-6 CAPLUS

CN 4-Piperidinepentanamide, N-methyl-1-[[tetrahydro-4-

[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-52-7 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[5-(1,3-dihydro-2H-isoindol-2-yl)-5-oxopentyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

RN 622388-53-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-(octahydro-2H-isoindol-2-yl)-5-oxopentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

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RN 622388-54-9 CAPLUS

CN 4-Piperidinepentanamide, 1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]-N-[4-(trifluoromethoxy)phenyl]- (9CI) (CA INDEX NAME)

$$F_3C-O$$

$$NH-C-(CH_2)_4$$

$$C-NH-OH$$

RN 622388-55-0 CAPLUS

CN 4-Piperidinepentanamide, N-methyl-N-(2-methylpropyl)-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-56-1 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-oxo-5-(1-piperidinyl)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-57-2 CAPLUS

CN 4-Piperidinepentanamide, N-(4-fluorophenyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-58-3 CAPLUS

CN 4-Piperidinepentanamide, N-methyl-N-phenyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-60-7 CAPLUS

CN 4-Piperidinepentanamide, N-cyclohexyl-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-61-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-oxo-5-(1-pyrrolidinyl)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-62-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-oxo-5-(4-phenyl-1-piperazinyl)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-63-0 CAPLUS

CN 4-Piperidinepentanamide, N-(2-furanylmethyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-67-4 CAPLUS

CN 4-Piperidinepentanoic acid, 1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]-, methyl ester (9CI) (CA INDEX NAME)

RN 622388-72-1 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[4-[(oxo-2-thienylacetyl)amino]butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-75-4 CAPLUS

CN 4-Piperidinebutanamide, N-(cyclohexylmethyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-76-5 CAPLUS

CN 4-Piperidinepentanamide, N-(cyclohexylmethyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-79-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[4-[[(1-phenylcyclopropyl)carbonyl]amino]butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-05-3 CAPLUS

CN 4-Piperidinebutanamide, N-(2-furanylmethyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-06-4 CAPLUS

CN 4-Piperidinebutanamide, N-methyl-N-(4-methylphenyl)-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-07-5 CAPLUS

CN 4-Piperidinebutanamide, N-methyl-N-phenyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-08-6 CAPLUS

CN 4-Piperidinebutanamide, N-cyclohexyl-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-09-7 CAPLUS

CN 4-Piperidinebutanamide, N-phenyl-N-propyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} Ph & O \\ \hline & Ph & O \\ \hline & & \\ n-Pr-N-C- (CH_2)_3 \end{array}$$

RN 622389-10-0 CAPLUS

CN 4-Piperidinebutanamide, N-(4-fluorophenyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-11-1 CAPLUS

CN 4-Piperidinebutanamide, N-ethyl-N-phenyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-12-2 CAPLUS

CN 4-Piperidinebutanoic acid, 1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]-, methyl ester (9CI) (CA INDEX NAME)

$$\begin{array}{c|c}
0 & & & \\
N & S & & \\
N & S & & \\
0 & C - NH - OH
\end{array}$$
MeO-C- (CH₂) 3

RN 622389-24-6 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[3-[3-[4-(trifluoromethoxy)phenyl]-1,2,4-oxadiazol-5-yl]propyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3C-O$$
 N
 $CH_2)_3$
 N
 $C-NH-OH$

RN 622389-27-9 CAPLUS

CN 4-Piperidinepentanamide, N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]-N-[4-(trifluoromethoxy)phenyl]- (9CI) (CA INDEX NAME)

$$F_3C-O$$
 Me
 O
 $N-C-(CH_2)_4$
 O
 $C-NH-OH$
 O

RN 622389-50-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(2-phenylethyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-73-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-{(1E,3E)-5-(4-morpholinyl)-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 851771-04-5 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[5-(2,3-dihydro-1H-indol-1-yl)-5-oxopentyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c}
 & \circ & \circ & \circ \\
 & N - C - (CH_2)_4 & \circ & \circ & \circ \\
 & O & C - NH - OH \\
 & O & OH - OH
\end{array}$$

IT 851771-07-8P 851771-08-9P 851771-09-0P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(prepn. of piperidinyl- and piperazinyl-sulfonylmethyl hydroxamic acids and their use as matrix metalloproteinase inhibitors)

RN 851771-07-8 CAPLUS

CN 4-Piperidinepentanoic acid, 1-[[tetrahydro-4-[[[(tetrahydro-2H-pyran-2-yl)oxy]amino]carbonyl]-2H-pyran-4-yl]sulfonyl]-, ethyl ester (9CI) (CA INDEX NAME)

RN 851771-08-9 CAPLUS

CN 4-Piperidinepentanoic acid, 1-[[tetrahydro-4-[[[(tetrahydro-2H-pyran-2-yl)oxy]amino]carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 851771-09-0 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[5-(2,3-dihydro-1H-indol-1-yl)-5-oxopentyl]-1-piperidinyl]sulfonyl]tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

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L4
     ANSWER 2 OF 4 CAPLUS COPYRIGHT 2005 ACS on STN
AN
     2005:409509 CAPLUS
     142:463765
DN
     Preparation of piperidinyl- and piperazinylsulfonylmethyl hydroxamic acids
ΤI
     and their use as protease inhibitors
     Brown, David L.; Grapperhaus, Margaret L.; Kassab, Darren J.; Massa, Mark
IN
     A.; Mcdonald, Joseph J.; Mullins, Patrick B.; Rico, Joseph G.; Schmidt,
                                                  SANG IE
     Michelle A.
PA
     Pharmacia Corporation, USA
     PCT Int. Appl., 644 pp.
SO
     CODEN: PIXXD2
DT
     Patent
     English
LA
FAN.CNT 3
     PATENT NO.
                        KIND
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                                            APPLICATION NO.
                                                                   DATE
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PT
     WO 2005042521
                         A2
                                20050512
                                            WO 2004-US36666
                                                                   20041103
     WO 2005042521
                         A3
                                20050707
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
        NE, SN, TD, TG
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                                            US 2003-700202
                                                                   20031103
PRAI US 2003-700202
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    US 2002-375598P
                         Ρ
                                20020425
    US 2002-380713P
                         Ρ
                                20020515
    US 2002-392021P
                         Р
                                20020627
    US 2003-618288
                         A2
                                20030425
OS
    MARPAT 142:463765
GI
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$$A^{1}$$
 A^{2} N R ?

AB Title compds. I [A1-2 = H, alkyl, alkoxyalkyl, etc.; Rx = halo, CN, OH, NO2, etc.; E2 = CO, COO, OCO, amino, etc.; E3 = alkyl, alkenyl, alkynyl, etc.] are prepd. For instance, 4-[[4-(5-butylpyrazin-2-yl)piperazin-1-yl]sulfonyl]-N-(hydroxy)tetrahydro-2H-pyran-4-carboxamide.bul.2HCl (II) is prepd. in 8 steps from 1-(tert-butoxycarbonyl)piperazine, 2-chloropyrazine, butylmagnesium chloride, bis(2-bromoethyl)ether and O-(tetrahydro-2H-pyran-2-yl)hydroxyamine. II exhibits Ki = >10,000 nM for MMP-1, 1.52 nM for MMP-2, 0.696 nM for MMP-9, 1.82 nM for MMP-13 and 4290 nM for MMP-14. I are useful for the treatment of conditions assocd. with MMP activity and/or aggrecanase activity.

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IT
     622387-48-8P 622389-78-0P 622390-02-7P
     622390-05-0P 622390-07-2P 622390-55-0P
     622390-62-9P 622390-68-5P 622390-72-1P
     622390-78-7P 622390-81-2P 622390-88-9P
     622390-93-6P 622390-99-2P 622391-09-7P
     622391-15-5P 622391-31-5P 622391-35-9P
     622391-39-3P 622391-43-9P 622391-83-7P
     851771-02-3P 851771-07-8P 851771-08-9P
     851771-09-0P
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (intermediate; prepn. of piperidinyl-and piperazinyl-sulfonylmethyl
        hydroxamic acids and their use as matrix metalloproteinase inhibitors)
RN
     622387-48-8 CAPLUS
CN
     2H-Pyran-4-carboxamide, tetrahydro-4-[(4-octyl-1-piperidinyl)sulfonyl]-N-
     [(tetrahydro-2H-pyran-2-yl)oxy] - (9CI) (CA INDEX NAME)
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RN 622389-78-0 CAPLUS
CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-[(phenylmethoxy)methyl]-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622390-02-7 CAPLUS
CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-[(2,2,3,3-tetrafluoropropoxy)methyl]-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c}
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C \\
O \\
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\end{array}$$

$$\begin{array}{c|c}
CH_2 - O - CH_2 - CF_2 - CHF_2 \\
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O \\
O \\
O \\
O
\end{array}$$

RN 622390-05-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[[4-(trifluoromethyl)phenyl]methoxy]methyl]-1-piperidinyl]sulfonyl]-(9CI) (CA INDEX NAME)

RN 622390-07-2 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-(1Z)-1-octenyl-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622390-55-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[5-(2,2,2-trifluoroethoxy)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622390-62-9 CAPLUS
CN 2H-Pyran-4-carboxamide, 4-[[4-(5-ethoxypentyl)-1piperidinyl]sulfonyl]tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI)
(CA INDEX NAME)

RN 622390-68-5 CAPLUS
CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-(4-propoxybutyl)-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622390-72-1 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[4-(4,4,4-trifluorobutoxy)butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622390-78-7 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(hexyloxy)methyl]-1-piperidinyl]sulfonyl]tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622390-81-2 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[(4,4,4-trifluorobutoxy)methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622390-88-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-(6,6,6-trifluorohexyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622390-93-6 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-(2-butoxyethyl)-1-piperidinyl]sulfonyl]tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622390-99-2 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[2-(4,4,4-trifluorobutoxy)ethyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622391-09-7 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[4-(3,3,3-trifluoropropoxy)butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622391-15-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-[2-(pentyloxy)ethyl]-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622391-31-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-(7,7,8,8,8-pentafluorooctyl)-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622391-35-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-y1)oxy]-4-[[4-(8,8,8-trifluoroocty1)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622391-39-3 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-(5,5,6,6,6-pentafluorohexyl)-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622391-43-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-(6,6,7,7,7-pentafluoroheptyl)-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622391-83-7 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[3-[4-(trifluoromethoxy)phenyl]-1,2,4-oxadiazol-5-yl]butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 851771-02-3 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-(1E)-1-octenyl-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 851771-07-8 CAPLUS

CN 4-Piperidinepentanoic acid, 1-[[tetrahydro-4-[[[(tetrahydro-2H-pyran-2-yl)oxy]amino]carbonyl]-2H-pyran-4-yl]sulfonyl]-, ethyl ester (9CI) (CA INDEX NAME)

RN 851771-08-9 CAPLUS

CN 4-Piperidinepentanoic acid, 1-[[tetrahydro-4-[[[(tetrahydro-2H-pyran-2-yl)oxy]amino]carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 851771-09-0 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[5-(2,3-dihydro-1H-indol-1-yl)-5-oxopentyl]-1-piperidinyl]sulfonyl]tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

IT 622382-02-9P 622385-57-3P 622385-59-5P

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622385-60-8P 622385-61-9P 622385-62-0P
622385-88-0P 622385-89-1P 622385-90-4P
622385-91-5P 622385-92-6P 622387-62-6P
622387-72-8P 622387-73-9P 622387-74-0P
622387-75-1P 622387-76-2P 622387-77-3P
622387-78-4P 622387-79-5P 622387-80-8P
622387-81-9P 622387-82-0P 622387-83-1P
622387-84-2P 622387-92-2P 622387-94-4P
622387-96-6P 622387-98-8P 622388-18-5P
622388-35-6P 622388-36-7P 622388-37-8P
622388-38-9P 622388-40-3P 622388-42-5P
622388-44-7P 622388-46-9P 622388-47-0P
622388-48-1P 622388-50-5P 622388-51-6P
622388-52-7P 622388-53-8P 622388-54-9P
622388-55-0P 622388-56-1P 622388-57-2P
622388-58-3P 622388-60-7P 622388-61-8P
622388-62-9P 622388-63-0P 622388-67-4P
622388-72-1P 622388-75-4P 622388-76-5P
622388-79-8P 622389-05-3P 622389-06-4P
622389-07-5P 622389-08-6P 622389-09-7P
622389-10-0P 622389-11-1P 622389-12-2P
622389-24-6P 622389-27-9P 622389-50-8P
622389-73-5P 851771-04-5P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
(Uses)
   (prepn. of piperidinyl-and piperazinyl-sulfonylmethyl hydroxamic acids
   and their use as matrix metalloproteinase inhibitors)
622382-02-9 CAPLUS
2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[(4-octyl-1-
piperidinyl)sulfonyl] - (9CI) (CA INDEX NAME)
```

RN 622385-57-3 CAPLUS
CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(phenylmethoxy)methyl]1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622385-59-5 CAPLUS
CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(2,2,3,3-tetrafluoropropoxy)methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN

CN

RN 622385-60-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(2,2,3,3,3-pentafluoropropoxy)methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3C-CF_2-CH_2-O-CH_2$$

$$0$$

$$0$$

$$0$$

$$0$$

$$0$$

$$C-NH-OH$$

$$0$$

RN 622385-61-9 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(2,2,3,3,4,4,4-heptafluorobutoxy)methyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

$$F_3C-CF_2-CF_2-CH_2-O-CH_2$$
 $N-S$
 $C-NH-OH$

RN 622385-62-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[[(2,2,3,3,4,4,5,5-octafluoropentyl)oxy]methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_2CH-(CF_2)_3-CH_2-O-CH_2$$
 $N-S$
 $C-NH-OH$

RN 622385-88-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(1Z)-1-octenyl-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622385-89-1 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(1E)-1-octenyl-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

Me (CH₂)
$$\frac{E}{5}$$

RN 622385-90-4 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[[[4-(trifluoromethyl)phenyl]methoxy]methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622385-91-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[[[4-(trifluoromethoxy)phenyl]methoxy]methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622385-92-6 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[[4-[[4-[(trifluoromethyl)thio]phenyl]methoxy]methyl]-1-piperidinyl]sulfonyl]-

(9CI) (CA INDEX NAME)

RN 622387-62-6 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-(2,2,2-trifluoroethoxy)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-72-8 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-(3-butoxypropyl)-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

$$n-BuO-(CH_2)_3$$
 $N-S$
 O
 $C-NH-OH$

RN 622387-73-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[3-(4,4,4-trifluorobutoxy)propyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-74-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-(3,3,3-trifluoropropoxy)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-75-1 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-(5-ethoxypentyl)-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

RN 622387-76-2 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(4-propoxybutyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-77-3 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[4-(4,4,4-trifluorobutoxy)butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3C-(CH_2)_3-O-(CH_2)_4$$
 N
 S
 $C-NH-OH$

RN 622387-78-4 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(hexyloxy)methyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

RN 622387-79-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(4,4,4-trifluorobutoxy)methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-80-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(6,6,6-trifluorohexyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-81-9 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-(2-butoxyethyl)-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

RN 622387-82-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[2-(4,4,4-trifluorobutoxy)ethyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-83-1 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[4-(3,3,3-trifluoropropoxy)butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-84-2 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[2-(pentyloxy)ethyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$Me^{-(CH_2)_4-0-CH_2-CH_2} = 0$$

$$0$$

$$0$$

$$0$$

$$0$$

$$0$$

$$0$$

$$0$$

$$0$$

RN 622387-92-2 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(7,7,8,8,8-pentafluorooctyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3C-CF_2-(CH_2)_6$$
 $N-S$
 $C-NH-OH$

RN 622387-94-4 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(8,8,8-trifluorooctyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-96-6 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(5,5,6,6,6-pentafluorohexyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3$$
C- CF_2 - $(CH_2)_4$
 0
 C
 C
 C
 C
 C
 C
 C
 C

RN 622387-98-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(6,6,7,7,7-pentafluoroheptyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3C-CF_2-(CH_2)_5$$
 $N-S$
 $C-NH-OH$

RN 622388-18-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[4-[3-[4-(trifluoromethoxy)phenyl]-1,2,4-oxadiazol-5-yl]butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-35-6 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[5-(3,4-dihydro-2(1H)-isoquinolinyl)-5-oxopentyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

RN 622388-36-7 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(1E,3E)-5-(3,4-dihydro-2(1H)-isoquinolinyl)-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-37-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(1E,3E)-5-oxo-5-(4-phenyl-1-piperazinyl)-1,3-pentadienyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-38-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(1E,3E)-5-[methyl(2-methylpropyl)amino]-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-40-3 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(1E,3E)-5-oxo-5-(1-piperidinyl)-1,3-pentadienyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-42-5 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(1E,3E)-5-(cyclohexylmethylamino)-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-44-7 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(1E,3E)-5-(2,3-dihydro-1H-indol-1-yl)-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-46-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(1E,3E)-5-[(3aR,7aS)-octahydro-2H-isoindol-2-yl]-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]-, rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

Double bond geometry as shown.

$$\begin{array}{c|c}
H & & \\
R & & \\
\end{array}$$

$$\begin{array}{c|c}
E & & \\
\end{array}$$

$$\begin{array}{c|c}
H & \\
\end{array}$$

RN 622388-47-0 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(1E,3E)-5-(1,3-dihydro-2H-isoindol-2-yl)-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-48-1 CAPLUS

CN 4-Piperidinepentanamide, N-methyl-N-2-propynyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-50-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-(4-morpholinyl)-5-oxopentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-51-6 CAPLUS

CN 4-Piperidinepentanamide, N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-52-7 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[5-(1,3-dihydro-2H-isoindol-2-yl)-5-oxopentyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

RN 622388-53-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-(octahydro-2H-isoindol-2-yl)-5-oxopentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-54-9 CAPLUS

CN 4-Piperidinepentanamide, 1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]-N-[4-(trifluoromethoxy)phenyl]- (9CI) (CA INDEX NAME)

RN 622388-55-0 CAPLUS

CN 4-Piperidinepentanamide, N-methyl-N-(2-methylpropyl)-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-56-1 CAPLUS

•CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-oxo-5-(1-piperidinyl)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-57-2 CAPLUS

CN 4-Piperidinepentanamide, N-(4-fluorophenyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-58-3 CAPLUS

CN 4-Piperidinepentanamide, N-methyl-N-phenyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} Ph & O \\ \hline | & \\ | & \\ Me-N-C- (CH_2) & 4 \end{array}$$

RN 622388-60-7 CAPLUS

CN 4-Piperidinepentanamide, N-cyclohexyl-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-61-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-oxo-5-(1-pyrrolidinyl)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-62-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-oxo-5-(4-phenyl-1-piperazinyl)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-63-0 CAPLUS

CN 4-Piperidinepentanamide, N-(2-furanylmethyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-67-4 CAPLUS

CN 4-Piperidinepentanoic acid, 1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]-, methyl ester (9CI) (CA INDEX NAME)

RN 622388-72-1 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[4-[(oxo-2-thienylacetyl)amino]butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-75-4 CAPLUS

CN 4-Piperidinebutanamide, N-(cyclohexylmethyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-76-5 CAPLUS

CN 4-Piperidinepentanamide, N-(cyclohexylmethyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-79-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[4-[[(1-phenylcyclopropyl)carbonyl]amino]butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-05-3 CAPLUS

CN 4-Piperidinebutanamide, N-(2-furanylmethyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-06-4 CAPLUS

CN 4-Piperidinebutanamide, N-methyl-N-(4-methylphenyl)-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-07-5 CAPLUS

CN 4-Piperidinebutanamide, N-methyl-N-phenyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-08-6 CAPLUS

CN 4-Piperidinebutanamide, N-cyclohexyl-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-09-7 CAPLUS

CN 4-Piperidinebutanamide, N-phenyl-N-propyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c}
\text{Ph O} & & & \\
\mid & \mid \\
\text{n-Pr-N-C-} (CH_2)_3 & & & \\
\end{array}$$

RN 622389-10-0 CAPLUS

CN 4-Piperidinebutanamide, N-(4-fluorophenyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

F Me O N C C
$$(CH_2)_3$$
 $N = S$ $C = NH = OH$ O

RN 622389-11-1 CAPLUS

CN 4-Piperidinebutanamide, N-ethyl-N-phenyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-12-2 CAPLUS

CN 4-Piperidinebutanoic acid, 1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]-, methyl ester (9CI) (CA INDEX NAME)

RN 622389-24-6 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[3-[3-[4-(trifluoromethoxy)phenyl]-1,2,4-oxadiazol-5-yl]propyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3C-O$$
 N
 $C-NH-OH$
 $C-NH-OH$

RN 622389-27-9 CAPLUS

CN 4-Piperidinepentanamide, N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]-N-[4-(trifluoromethoxy)phenyl]- (9CI) (CA INDEX NAME)

RN 622389-50-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(2-phenylethyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & & & & & \\ & & & & & & \\ \text{Ph-} & \text{CH}_2 - \text{CH}_2 & & & & \\ & & & & & & \\ \end{array}$$

RN 622389-73-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(1E,3E)-5-(4-morpholinyl)-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 851771-04-5 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[5-(2,3-dihydro-1H-indol-1-yl)-5-oxopentyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

L4 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2005 ACS on STN

AN 2003:875282 CAPLUS

DN 139:364961

TI Preparation of piperidinyl-and piperazinyl-sulfonylmethyl hydroxamic acids and their use as protease inhibitors

Barta, Thomas E.; Becker, Daniel P.; Bedell, Louis J.; Boehm, Terri L.; Brown, David L.; Carroll, Jeffery N.; Chen, Yiyuan; Fobian, Yvette; Freskos, John N.; Gasiecki, Alan F.; Grapperhaus, Margaret; Heintz, Robert M.; Hockerman, Susan L.; Kassab, Darren J.; Khanna, Ish Kumar; Kolodziej, Stephen A.; Massa, Mark; Mcdonald, Joseph; Mischke, Brent V.; Mischke, Deborah A.; Mullins, Patrick B.; Nagy, Mark; Norton, Monica B.; Rico, Joseph G.; Schmidt, Michelle A.; Stehle, Nathan W.; Talley, John J.; Vernier, William F.; Villamill, Clara I.; Wang, Lijuan Jane; Wynn, Thomas

PA Pharmacia Corporation, USA; et al.

SO PCT Int. Appl., 819 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 3

| | PATENT NO. | | | | | KIND | | DATE | | APPLICATION NO. | | | | | | DATE | | |
|----|---------------|-----|-----|-----|-----|------|----------|------|-----------------|-----------------|-----|-----|-----|-----|---|------|-----|-----|
| | | | | | | | | | | | | | | | | | | |
| ΡI | WO 2003091247 | | | | A2 | | 20031106 | | WO 2003-US13123 | | | | | | 20030425 | | | |
| | WO 2003091247 | | | | A3 | | 20040115 | | | | | | | | • | | | |
| | | W: | ΑE, | AG, | ΑL, | AM, | ΑT, | AU, | ΑZ, | BA, | BB, | BG, | BR, | BY, | BZ, | CA, | CH, | CN, |
| | | | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DZ, | EC, | EE, | ES, | FI, | GB, | GD. | GE. | GH. |
| | | | | | | | | IN, | | | | | | | | | | |
| | | | | | | | | MD, | | | | | | | | | | |
| | | | | | | | | SC, | | | | | | | | | | |
| | | | | | | | | VC, | | | | | | | | | • | • |
| | | RW: | GH, | GM, | ΚE, | LS, | MW, | MZ, | SD, | SL, | SZ, | TZ, | ŪĠ, | ZM, | ZW, | AM, | AZ, | BY, |
| | | | KG, | KZ, | MD, | RU, | TJ, | TM, | ΑT, | BE, | BG, | CH, | CY, | CZ, | DE, | DK, | EE, | ES, |
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BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
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                                             CA 2003-2483314
                                                                     20030425
     EP 1501827
                          A2
                                 20050202
                                             EP 2003-718529
                                                                     20030425
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
     BR 2003009671
                          Α
                               . 20050503
                                             BR 2003-9671
                                                                     20030425
PRAI US 2002-375598P
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                                20020425
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     US 2002-380713P
                                20020515
                          P
     US 2002-392021P
                                 20020627
     WO 2003-US13123
                          W
                                 20030425
os
     MARPAT 139:364961
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* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

Title compds. I [A1 and A2 together with the C to which they are bonded AB join to form (un) substituted-heterocyclyl or -carbocyclyl, or A1 and A2 are independently selected from H, alkyl, alkoxyalkyl, alkenyl, alkynyl, etc.; Rx = H, halo, CN, OH, NO2, alkyl, alkenyl, alkoxy, alkoxyalkyl, heterocyclyl, etc.; Y = N, CH, or CRx; E1 = (un) substituted heteroaryl; E2 = 0, CO, C(0)0, OC(0), bond, S, etc.; E3 = halo, CN, (un)substitutedalkyl, -alkenyl, -alkynyl, -heterocyclyl, heterocyclylalkyl, etc.] and their pharmaceutically acceptable salts are prepd. and disclosed as protease inhibitors. Thus, e.g., II.cntdot.HCl was prepd. with piperazine ring formation occurring via cyclization of 2,2,2-trifluoroethoxyaniline (prepn. given) with N, N-di(2-chloroethyl) methylsulfonamide (prepn. given) to provide piperazinyl intermediate III which was converted in five addnl. steps to the desired product. This invention is directed generally to proteinase (also known as 'protease') inhibitors, and more particularly, inhibitors of matrix metalloproteinase (also known as 'matrix metalloprotease' or 'MMP') activity and/or aggrecanase activity. assays to det. inhibition consts. (Ki) against MMP-1, MMP-2, MMP-9, MMP-13 and MMP-14, I possessed values ranging from 0.13->10,000. This invention also is directed to compns. of such hydroxamic acids, intermediates for the syntheses of such hydroxamic acids, methods for making such hydroxamic acids, and methods for treating conditions (particularly pathol. conditions) assocd. with MMP activity and/or aggrecanase activity.

IT 622396-56-9P 622396-57-0P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(claimed compds.; prepn. of piperidinyl-and piperazinyl-sulfonylmethyl hydroxamic acids and their use as matrix metalloproteinase inhibitors) 622396-56-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(1E,3E)-5-(methyl-2-propynylamino)-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]- (9CI) (CAINDEX NAME)

Double bond geometry as shown.

RN

RN 622396-57-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[3-(pentyloxy)propyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$Me^{-(CH_2)_4-O^{-(CH_2)_3}}$$

ΙT 622387-48-8P 622389-78-0P 622390-02-7P 622390-05-0P 622390-07-2P 622390-55-0P 622390-62-9P 622390-68-5P 622390-72-1P 622390-78-7P 622390-81-2P 622390-88-9P 622390-93-6P 622390-99-2P 622391-09-7P 622391-15-5P 622391-31-5P 622391-35-9P 622391-39-3P 622391-43-9P 622391-83-7P RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (intermediate; prepn. of piperidinyl-and piperazinyl-sulfonylmethyl hydroxamic acids and their use as matrix metalloproteinase inhibitors) RN 622387-48-8 CAPLUS CN 2H-Pyran-4-carboxamide, tetrahydro-4-[(4-octyl-1-piperidinyl)sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622389-78-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-[(phenylmethoxy)methyl]-1-

piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622390-02-7 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-[(2,2,3,3-tetrafluoropropoxy)methyl]-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622390-05-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[[4-(trifluoromethyl)phenyl]methoxy]methyl]-1-piperidinyl]sulfonyl](9CI) (CA INDEX NAME)

RN 622390-07-2 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-(1Z)-1-octenyl-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622390-55-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[5-(2,2,2-trifluoroethoxy)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622390-62-9 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-(5-ethoxypentyl)-1-

piperidinyl]sulfonyl]tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI)
(CA INDEX NAME)

RN 622390-68-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-(4-propoxybutyl)-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622390-72-1 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[4-(4,4,4-trifluorobutoxy)butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622390-78-7 CAPLUS
CN 2H-Pyran-4-carboxamide, 4-[[4-[(hexyloxy)methyl]-1piperidinyl]sulfonyl]tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI)
(CA INDEX NAME)

RN 622390-81-2 CAPLUS
CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4[(4,4,4-trifluorobutoxy)methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622390-88-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-(6,6,6-trifluorohexyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622390-93-6 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-(2-butoxyethyl)-1-piperidinyl]sulfonyl]tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622390-99-2 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[2-(4,4,4-trifluorobutoxy)ethyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622391-09-7 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[4-(3,3,3-trifluoropropoxy)butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622391-15-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-[2-(pentyloxy)ethyl]-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

$$O = C O C O CH_2 - CH_2 - O - (CH_2)_4 - Me$$

$$O = C O CH_2 - CH_2 - O - (CH_2)_4 - Me$$

RN 622391-31-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-(7,7,8,8,8-pentafluorooctyl)-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622391-35-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-(8,8,8-trifluorooctyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622391-39-3 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-(5,5,6,6,6-pentafluorohexyl)-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622391-43-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-(6,6,7,7,7-pentafluoroheptyl)-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 622391-83-7 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]-4-[[4-[3-[4-(trifluoromethoxy)phenyl]-1,2,4-oxadiazol-5-yl]butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

IT 622382-02-9P 622385-57-3P 622385-59-5P 622385-60-8P 622385-61-9P 622385-62-0P 622385-88-0P 622385-89-1P 622385-90-4P 622385-91-5P 622385-92-6P 622387-62-6P 622387-72-8P 622387-73-9P 622387-74-0P 622387-75-1P 622387-76-2P 622387-77-3P 622387-78-4P 622387-79-5P 622387-80-8P 622387-81-9P 622387-82-0P 622387-83-1P 622387-84-2P 622387-92-2P 622387-94-4P 622387-96-6P 622387-98-8P 622388-18-5P 622388-35-6P 622388-36-7P 622388-37-8P 622388-38-9P 622388-40-3P 622388-42-5P 622388-44-7P 622388-46-9P 622388-47-0P 622388-48-1P 622388-50-5P 622388-51-6P 622388-52-7P 622388-53-8P 622388-54-9P 622388-55-0P 622388-56-1P 622388-57-2P 622388-58-3P 622388-60-7P 622388-61-8P 622388-62-9P 622388-63-0P 622388-67-4P 622388-72-1P 622388-75-4P 622388-76-5P 622388-79-8P 622389-05-3P 622389-06-4P 622389-07-5P 622389-08-6P 622389-09-7P 622389-10-0P 622389-11-1P 622389-12-2P 622389-24-6P 622389-27-9P 622389-50-8P 622389-73-5P RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (prepn. of piperidinyl-and piperazinyl-sulfonylmethyl hydroxamic acids and their use as matrix metalloproteinase inhibitors) RN 622382-02-9 CAPLUS 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[(4-octyl-1-CN piperidinyl)sulfonyl]- (9CI) (CA INDEX NAME)

RN 622385-57-3 CAPLUS
CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(phenylmethoxy)methyl]1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622385-59-5 CAPLUS
CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(2,2,3,3-tetrafluoropropoxy)methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_2CH-CF_2-CH_2-O-CH_2$$

$$0$$

$$0$$

$$0$$

$$0$$

$$0$$

$$0$$

$$0$$

$$0$$

$$0$$

RN 622385-60-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(2,2,3,3,3-pentafluoropropoxy)methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3C-CF_2-CH_2-O-CH_2$$

$$0$$

$$0$$

$$C-NH-OH$$

$$0$$

RN 622385-61-9 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(2,2,3,3,4,4,4-heptafluorobutoxy)methyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

$$F_3C-CF_2-CF_2-CH_2-O-CH_2$$

$$0$$

$$0$$

$$C-NH-OH$$

$$0$$

RN 622385-62-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[[(2,2,3,3,4,4,5,5-octafluoropentyl)oxy]methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_2$$
CH- (CF₂)₃-CH₂-O-CH₂ O C -NH-OH

RN 622385-88-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(1Z)-1-octenyl-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622385-89-1 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(1E)-1-octenyl-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

Me (CH₂)
$$\frac{E}{5}$$

RN 622385-90-4 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[[4-(trifluoromethyl)phenyl]methoxy]methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$_{\text{CH}_2-\text{O-CH}_2}$$

RN 622385-91-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[[4-(trifluoromethoxy)phenyl]methoxy]methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3C-O$$
 CH_2-O-CH_2
 O
 $C-NH-OH$

RN 622385-92-6 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[[4-[[4-[(trifluoromethyl)thio]phenyl]methoxy]methyl]-1-piperidinyl]sulfonyl]-(9CI) (CA INDEX NAME)

RN 622387-62-6 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-(2,2,2-trifluoroethoxy)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-72-8 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-(3-butoxypropyl)-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

$$n-BuO-(CH_2)$$
 3 $N-S$ $C-NH-OH$

RN 622387-73-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[3-(4,4,4-trifluorobutoxy)propyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$_{\rm F_3C-\ (CH_2)\ 3}$$
 - 0- (CH₂) $_{\rm 3}$ - 0- (CH₂) $_{\rm 3}$ - 0- (CH₂) $_{\rm 3}$

RN 622387-74-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-(3,3,3-trifluoropropoxy)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-75-1 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-(5-ethoxypentyl)-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

RN 622387-76-2 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(4-propoxybutyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-77-3 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[4-(4,4,4-trifluorobutoxy)butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3C-(CH_2)_3-O-(CH_2)_4$$
 $N-S$
 $C-NH-OH$

RN 622387-78-4 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(hexyloxy)methyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

$$N = (CH_2)_5 - O - CH_2$$
 $N = (CH_2)_5 - O - CH_2$
 $N = (CH_2)_5 - O - CH_2$
 $N = (CH_2)_5 - O - CH_2$

RN 622387-79-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(4,4,4-trifluorobutoxy)methyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$_{F_3C- (CH_2)_3-O-CH_2}$$
 $N-S$
 0
 $C-NH-OH$
 0

RN 622387-80-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(6,6,6-trifluorohexyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-81-9 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-(2-butoxyethyl)-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

RN 622387-82-0 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[2-(4,4,4-trifluorobutoxy)ethyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-83-1 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[4-(3,3,3-trifluoropropoxy)butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$_{\text{F}_3\text{C}-\text{CH}_2-\text{CH}_2-\text{O}-\text{(CH}_2)}_{4}$$

RN 622387-84-2 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[2-(pentyloxy)ethyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622387-92-2 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(7,7,8,8,8-pentafluorooctyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3C-CF_2-(CH_2)_6$$
 $N-S$
 $C-NH-OH$

RN 622387-94-4 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(8,8,8-trifluorooctyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3C-(CH_2)_7$$

N

S

C

C

N

O

C

N

O

C

N

O

O

RN 622387-96-6 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(5,5,6,6,6-pentafluorohexyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3C-CF_2-(CH_2)$$

RN 622387-98-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(6,6,7,7,7-pentafluoroheptyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$F_3C-CF_2-(CH_2)_5$$
 $N-S$
 $C-NH-OH$

RN 622388-18-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[4-[3-[4-(trifluoromethoxy)phenyl]-1,2,4-oxadiazol-5-yl]butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-35-6 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[5-(3,4-dihydro-2(1H)-isoquinolinyl)-5-oxopentyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

RN 622388-36-7 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(1E,3E)-5-(3,4-dihydro-2(1H)-isoquinolinyl)-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-37-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(1E,3E)-5-oxo-5-(4-phenyl-1-piperazinyl)-1,3-pentadienyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-38-9 CAPLUS

Double bond geometry as shown.

RN 622388-40-3 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(1E,3E)-5-oxo-5-(1-piperidinyl)-1,3-pentadienyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-42-5 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(1E,3E)-5-(cyclohexylmethylamino)-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-44-7 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(1E,3E)-5-(2,3-dihydro-1H-indol-1-yl)-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-46-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(1E,3E)-5-[(3aR,7aS)-octahydro-2H-isoindol-2-yl]-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]-, rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

Double bond geometry as shown.

RN 622388-47-0 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(1E,3E)-5-(1,3-dihydro-2H-isoindol-2-yl)-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

RN 622388-48-1 CAPLUS

CN 4-Piperidinepentanamide, N-methyl-N-2-propynyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-50-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-(4-morpholinyl)-5-oxopentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

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RN 622388-51-6 CAPLUS

CN 4-Piperidinepentanamide, N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-52-7 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[5-(1,3-dihydro-2H-isoindol-2-yl)-5-oxopentyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy- (9CI) (CA INDEX NAME)

RN 622388-53-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-(octahydro-2H-isoindol-2-yl)-5-oxopentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-54-9 CAPLUS

CN 4-Piperidinepentanamide, 1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2Hpyran-4-yl]sulfonyl]-N-[4-(trifluoromethoxy)phenyl]- (9CI) (CA INDEX
NAME)

RN 622388-55-0 CAPLUS

CN 4-Piperidinepentanamide, N-methyl-N-(2-methylpropyl)-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-56-1 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-oxo-5-(1-piperidinyl)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-57-2 CAPLUS

CN 4-Piperidinepentanamide, N-(4-fluorophenyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-58-3 CAPLUS

CN 4-Piperidinepentanamide, N-methyl-N-phenyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-60-7 CAPLUS

CN 4-Piperidinepentanamide, N-cyclohexyl-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-61-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-oxo-5-(1-pyrrolidinyl)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-62-9 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[5-oxo-5-(4-phenyl-1-piperazinyl)pentyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-63-0 CAPLUS

CN 4-Piperidinepentanamide, N-(2-furanylmethyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-67-4 CAPLUS

CN 4-Piperidinepentanoic acid, 1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]-, methyl ester (9CI) (CA INDEX NAME)

RN 622388-72-1 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[4-[4-[(oxo-2-thienylacetyl)amino]butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-75-4 CAPLUS

CN 4-Piperidinebutanamide, N-(cyclohexylmethyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-76-5 CAPLUS

CN 4-Piperidinepentanamide, N-(cyclohexylmethyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622388-79-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[4-[[(1-phenylcyclopropyl)carbonyl]amino]butyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-05-3 CAPLUS

CN 4-Piperidinebutanamide, N-(2-furanylmethyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-06-4 CAPLUS

CN 4-Piperidinebutanamide, N-methyl-N-(4-methylphenyl)-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-07-5 CAPLUS

CN 4-Piperidinebutanamide, N-methyl-N-phenyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} Ph & O \\ \parallel & \parallel \\ Me-N-C-(CH_2) & 3 \end{array}$$

RN 622389-08-6 CAPLUS

CN 4-Piperidinebutanamide, N-cyclohexyl-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-09-7 CAPLUS

CN 4-Piperidinebutanamide, N-phenyl-N-propyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

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RN 622389-10-0 CAPLUS

CN 4-Piperidinebutanamide, N-(4-fluorophenyl)-N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-11-1 CAPLUS

CN 4-Piperidinebutanamide, N-ethyl-N-phenyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-12-2 CAPLUS

CN 4-Piperidinebutanoic acid, 1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]-, methyl ester (9CI) (CA INDEX NAME)

RN 622389-24-6 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[3-[3-[4-(trifluoromethoxy)phenyl]-1,2,4-oxadiazol-5-yl]propyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

RN 622389-27-9 CAPLUS

CN 4-Piperidinepentanamide, N-methyl-1-[[tetrahydro-4-[(hydroxyamino)carbonyl]-2H-pyran-4-yl]sulfonyl]-N-[4-(trifluoromethoxy)phenyl]- (9CI) (CA INDEX NAME)

RN 622389-50-8 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(2-phenylethyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & & \\ \text{Ph-} & \text{CH}_2 - \text{CH}_2 \end{array} \\ \end{array}$$

RN 622389-73-5 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-[(1E,3E)-5-(4-morpholinyl)-5-oxo-1,3-pentadienyl]-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

L4

ANSWER 4 OF 4 CAPLUS COPYRIGHT 2005 ACS on STN

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AN
      2000:553575 CAPLUS
DN
      133:164006
      Preparation of sulfamato hydroxamic acid metalloprotease inhibitors
TI
     De Crescenzo, Gary A.; Rico, Joseph G.; Boehm, Terri L.; Carroll, Jeffery
IN
     N.; Kassab, Darren J.; Mischke, Deborah A.
PA
      G.D. Searle and Co., USA
      PCT Int. Appl., 628 pp.
so
      CODEN: PIXXD2 .
DT
      Patent
      English
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FAN.CNT 1
      PATENT NO.
                             KIND
                                     DATE
                                                   APPLICATION NO.
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      WO 2000046221
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                                                   WO 2000-US3061
                              A1
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               IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM,
               AZ, BY, KG, KZ, MD, RU, TJ, TM
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                                                CA 2000-2362230
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     MARPAT 133:164006
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HO
$$SO_2 - N$$
 $O - CH_2 - Ph$

AΒ The title compds. R2OC(O)CR1R2SO2NR3aR3b (I) [wherein R1 and R2 taken together with the C to which they are attached = (un)substituted heterocyclyl or cycloalkyl; or R1 and R2 = independently H, (un) substituted (cyclo) alkyl, alkyloxylalkyl, alkylthioalkyl, alkenyl, alkynyl, aryl(alkyl), heterocyclyl(alkyl), etc.; R3a and R3b = independently H or (un) substituted alkyl, alkenyl, alkynyl, (hetero) aryl, heterocyclyl, cycloalkyl, or alkoxyalkyl; R20 = OH, alkoxyl, aryloxy, NH-OR22, or NH-OR14; R22 = selectively removable protecting group, such as 2-THP, benzyl, trisubstituted silyl, o-NO2C6H4, etc.; R14 = H, a cation, or acyl] were prepd. as selective matrix metalloproteinase (MMP) inhibitors for the treatment of various conditions, such as pathol. breakdown of connective tissue, osteoarthritis, inflammation, tumor growth, and angiogenesis. Examples include the syntheses of over 50 piperidinylsulfonyl and piperazinylsulfonyl hydroxamic acids and their intermediates. In vitro MMP assay data for I show selective inhibition of MMP-2 and MMP-13 compared to MMP-1. Some inhibition assay data for MMP-3, MMP-7, MMP-8, MMP-9, and MMP-14 are also given. Thus, II was prepd. in a multi-step sequence involving addn. of MeOC(0)Cl to 1-(methylsulfonyl)-4-(benzyloxy)piperidine (4-step prepn. given) to form the methylene sulfonamide, cycloaddn. of dibromodiethyl ether to give the THF-substituted sulfonamide, deesterification, addn. of O-(tetrahydro-2H-pyran-2-yl)hydroxylamine to form the THP hydroxamate, and deprotection to yield the desired hydroxamic acid. II inhibited MMP-1, MMP-2, and MMP-13 with IC50 values of < 10,000 nM, 7.0 nM and 20.0 nM,

II

IT 287952-41-4P 287954-44-3P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(intermediate; prepn. of sulfamato hydroxamic acid metalloprotease inhibitors by cycloaddn. of dihalodialkyl ethers and amines to methylene sulfonamides followed by addn. of hydroxylamines)

RN 287952-41-4 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-4-[[4-(phenylmethyl)-1-piperidinyl]sulfonyl]-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

RN 287954-44-3 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[(3,5-dimethyl-1-piperidinyl)carbonyl]-1-piperidinyl]sulfonyl]tetrahydro-N-[(tetrahydro-2H-pyran-2-yl)oxy]- (9CI) (CA INDEX NAME)

IT 287951-55-7P 287951-75-1P 287951-95-5P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (target compd.; prepn. of sulfamato hydroxamic acid metalloprotease inhibitors by cycloaddn. of dihalodialkyl ethers and amines to methylene sulfonamides followed by addn. of hydroxylamines)

RN 287951-55-7 CAPLUS

CN 2H-Pyran-4-carboxamide, tetrahydro-N-hydroxy-4-[[4-(phenylmethyl)-1-piperidinyl]sulfonyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c}
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 & N & S & O \\
 & O & C - NH - OH \\
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 & O & O &$$

RN 287951-75-1 CAPLUS

10/700202 .

CN 2H-Pyran-4-carboxamide, 4-[[4-[[(3R,5S)-3,5-dimethyl-1piperidinyl]carbonyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy-, rel(9CI) (CA INDEX NAME)

Relative stereochemistry.

RN 287951-95-5 CAPLUS

CN 2H-Pyran-4-carboxamide, 4-[[4-[[(3R,5R)-3,5-dimethyl-1piperidinyl]carbonyl]-1-piperidinyl]sulfonyl]tetrahydro-N-hydroxy-, rel(9CI) (CA INDEX NAME)

Relative stereochemistry.

RE.CNT 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> file caold COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 20.21 182.18 DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION CA SUBSCRIBER PRICE -2.92 -2.92

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printed between 1907-1966 are available in the PAGE display formats.

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